

RAW SEQUENCE LISTING

**The Biotechnology Systems Branch of the Scientific and Technical
Information Center (STIC) no errors detected.**

Application Serial Number: 10/562,942
Source: IFJP
Date Processed by STIC: 1/10/06

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IFWP

RAW SEQUENCE LISTING

DATE: 01/10/2006

PATENT APPLICATION: US/10/562,942

TIME: 08:53:10

Input Set : A:\PTO.DA.txt

Output Set: N:\CRF4\01102006\J562942.raw

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3 <110> APPLICANT: DOI, Hirofumi
4      SAITO, Ken
6 <120> TITLE OF INVENTION: Inhibition of Nerve Cell Death by Inhibiting Degradation of
SHC3, ATF6 or
7      CREBL1 by HtrA2 and Method of Ameliorating Neurodegenerative Diseases
9 <130> FILE REFERENCE: 3190-088
C--> 11 <140> CURRENT APPLICATION NUMBER: US/10/562,942
12 <141> CURRENT FILING DATE: 2005-12-29
14 <150> PRIOR APPLICATION NUMBER: PCT/JP2004/014378
15 <151> PRIOR FILING DATE: 2004-09-30
17 <150> PRIOR APPLICATION NUMBER: JP P2003-342588
18 <151> PRIOR FILING DATE: 2003-09-30
20 <160> NUMBER OF SEQ ID NOS: 37
22 <170> SOFTWARE: PatentIn version 3.1
24 <210> SEQ ID NO: 1
25 <211> LENGTH: 1377
26 <212> TYPE: DNA
27 <213> ORGANISM: Homo sapiens
29 <220> FEATURE:
30 <221> NAME/KEY: misc_feature
31 <223> OTHER INFORMATION: DNA that codes for HtrA2 precursor protein
34 <400> SEQUENCE: 1
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37 gggggcattc gctgggggag gagaccccggt ttgacccttg acctccgggc cctgctgacg      120
39 tcaggaactt ctgacccccg gggccgagtg acttatggga ccccgagctt ctgggcccgg      180
41 ttgtctgttg gggtcactga accccgagca tgcctgacgt ctgggacccc ggggtccccg      240
43 gcacaactga ctgcggtgac ccagataacc aggaccggg aggcctcaga gaactctgga      300
45 acccgttcgc gcgcgtggct ggcggtggcg ctgggcgctg ggggggcagt gctgttggtg      360
47 ttgtggggcg ggggtcgggg tcctccggcc gtctcgcg cgtccctag cccgcccgcc      420
49 gcttctcccc ggagtcagta caacttcac gcagatgtgg tggagaagac agcacctgcc      480
51 gtgggtctata tcgagatcct ggaccggcac ctttctctgg gccgcgaggt ccctatctcg      540
53 aacggctcag gattcgtggt ggctgccgat gggctcattg tcaccaacgc ccatgtggtg      600
55 gctgatcggc gcagagtcgg tgtgagactg ctaagcggcg acacgtatga ggccgtggtc      660
57 acagctgtgg atcccgtggc agacatcgca acgctgagga ttcagactaa ggagcctctc      720
59 cccacgctgc ctctgggacg ctgagctgat gtccggcaag gggagtttgt tgttgccatg      780
61 ggaagtcctt ttgactgca gaacacgatc acatccggca ttgttagctc tgctcagcgt      840
63 ccagccagag acctgggact cccccaaacc aatgtggaat acattcaaac tgatgcagct      900
65 attgattttg gaaactctgg aggtcccctg gttaacctgg atggggaggt gattggagtg      960
67 aacaccatga aggtcacagc tggaaatctc tttgccatcc cttctgatcg tcttcgagag      1020
69 tttctgcatc gtggggaaaa gaagaattcc tcctccggaa tcagtgggtc ccagcggcgc      1080
71 tacattgggg tgatgatgct gaccctgagt cccagcatcc ttgctgaact acagcttcga      1140
73 gaaccaagct ttcccgatgt tcagctgagt gtactatccc ataaagtcac cctgggctcc      1200
75 cctgcacacc gggctgggtc gcggcctggg gatgtgattt tggccattgg ggagcagatg      1260
77 gtacaaaatg ctgaagatgt ttatgaagct gttcgaacct aatcccagtt ggcagtgcag      1320

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79 atccggcgagg gacgagaaac actgacctta tatgtgaccc ctgaggtcac agaatga      1377
82 <210> SEQ ID NO: 2
83 <211> LENGTH: 458
84 <212> TYPE: PRT
85 <213> ORGANISM: Homo sapiens
87 <220> FEATURE:
88 <221> NAME/KEY: misc_feature
89 <223> OTHER INFORMATION: HtrA2 precursor protein
92 <400> SEQUENCE: 2
94 Met Ala Ala Pro Arg Ala Gly Arg Gly Ala Gly Trp Ser Leu Arg Ala
95 1          5          10          15
98 Trp Arg Ala Leu Gly Gly Ile Arg Trp Gly Arg Arg Pro Arg Leu Thr
99          20          25          30
102 Pro Asp Leu Arg Ala Leu Leu Thr Ser Gly Thr Ser Asp Pro Arg Ala
103          35          40          45
106 Arg Val Thr Tyr Gly Thr Pro Ser Leu Trp Ala Arg Leu Ser Val Gly
107          50          55          60
110 Val Thr Glu Pro Arg Ala Cys Leu Thr Ser Gly Thr Pro Gly Pro Arg
111 65          70          75          80
114 Ala Gln Leu Thr Ala Val Thr Pro Asp Thr Arg Thr Arg Glu Ala Ser
115          85          90          95
118 Glu Asn Ser Gly Thr Arg Ser Arg Ala Trp Leu Ala Val Ala Leu Gly
119          100         105         110
122 Ala Gly Gly Ala Val Leu Leu Leu Leu Trp Gly Gly Gly Arg Gly Pro
123          115         120         125
127 Pro Ala Val Leu Ala Ala Val Pro Ser Pro Pro Pro Ala Ser Pro Arg
128          130         135         140
131 Ser Gln Tyr Asn Phe Ile Ala Asp Val Val Glu Lys Thr Ala Pro Ala
132 145         150         155         160
135 Val Val Tyr Ile Glu Ile Leu Asp Arg His Pro Phe Leu Gly Arg Glu
136          165         170         175
139 Val Pro Ile Ser Asn Gly Ser Gly Phe Val Val Ala Ala Asp Gly Leu
140          180         185         190
143 Ile Val Thr Asn Ala His Val Val Ala Asp Arg Arg Arg Val Arg Val
144          195         200         205
147 Arg Leu Leu Ser Gly Asp Thr Tyr Glu Ala Val Val Thr Ala Val Asp
148          210         215         220
151 Pro Val Ala Asp Ile Ala Thr Leu Arg Ile Gln Thr Lys Glu Pro Leu
152 225         230         235         240
155 Pro Thr Leu Pro Leu Gly Arg Ser Ala Asp Val Arg Gln Gly Glu Phe
156          245         250         255
159 Val Val Ala Met Gly Ser Pro Phe Ala Leu Gln Asn Thr Ile Thr Ser
160          260         265         270
163 Gly Ile Val Ser Ser Ala Gln Arg Pro Ala Arg Asp Leu Gly Leu Pro
164          275         280         285
167 Gln Thr Asn Val Glu Tyr Ile Gln Thr Asp Ala Ala Ile Asp Phe Gly
168          290         295         300
171 Asn Ser Gly Gly Pro Leu Val Asn Leu Asp Gly Glu Val Ile Gly Val
172 305         310         315         320

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```

175 Asn Thr Met Lys Val Thr Ala Gly Ile Ser Phe Ala Ile Pro Ser Asp
176          325          330          335
179 Arg Leu Arg Glu Phe Leu His Arg Gly Glu Lys Lys Asn Ser Ser Ser
180          340          345          350
183 Gly Ile Ser Gly Ser Gln Arg Arg Tyr Ile Gly Val Met Met Leu Thr
184          355          360          365
187 Leu Ser Pro Ser Ile Leu Ala Glu Leu Gln Leu Arg Glu Pro Ser Phe
188          370          375          380
191 Pro Asp Val Gln His Gly Val Leu Ile His Lys Val Ile Leu Gly Ser
192 385          390          395          400
195 Pro Ala His Arg Ala Gly Leu Arg Pro Gly Asp Val Ile Leu Ala Ile
196          405          410          415
199 Gly Glu Gln Met Val Gln Asn Ala Glu Asp Val Tyr Glu Ala Val Arg
200          420          425          430
203 Thr Gln Ser Gln Leu Ala Val Gln Ile Arg Arg Gly Arg Glu Thr Leu
204          435          440          445
207 Thr Leu Tyr Val Thr Pro Glu Val Thr Glu
208          450          455

```

211 <210> SEQ ID NO: 3

212 <211> LENGTH: 981

213 <212> TYPE: DNA

214 <213> ORGANISM: Homo sapiens

216 <220> FEATURE:

217 <221> NAME/KEY: misc_feature

218 <223> OTHER INFORMATION: DNA that codes for mature HtrA2

221 <400> SEQUENCE: 3

```

222 atggcgcgtcc ctagcccgcc gcccgcgttct ccccgaggatc agtacaactt catcgcagat      60
224 gtgggtggaga agacagcacc tgccgtgggtc tatatcgaga tcctggaccg gcaccctttc      120
226 ttgggcccgcg aggtccctat ctcgaaacggc tcaggattcg tgggtggctgc cgatgggctc      180
228 attgtcacca acgcccattgt ggtggctgat cggcgcagag tccgtgtgag actgctaagc      240
230 ggcgacacgt atgaggccgt ggtcacagct gtggatcccg tggcagacat cgcaacgctg      300
232 aggattcaga ctaaggagcc tctccccacg ctgcctctgg gacgctcagc tgatgtccgg      360
234 caaggggaggt ttgttgttgc catgggaagt ccctttgcac tgcagaacac gatcacatcc      420
236 ggcattgtta gctctgctca gcgtccagcc agagacctgg gactccccca aaccaatgtg      480
238 gaatacattc aaactgatgc agctattgat tttggaaact ctggagggtcc cctggttaac      540
240 ctggatgggg aggtgattgg agtgaacacc atgaagggtc cagctggaat ctctttgcc      600
242 atcccttctg atcgtcttcg agagtctctg catcgtgggg aaaagaagaa ttcctcctcc      660
244 ggaatcagtg ggtcccagcg gcgctacatt ggggtgatga tgctgaccct gagtcccagc      720
246 atccttgctg aactacagct tcgagaacca agctttcccg atgttcagca tgggtgtactc      780
248 atccataaag tcatcctggg ctccccctgca caccgggctg gtctgcgggc tgggtgatgtg      840
250 attttggcca ttggggagca gatggtacaa aatgctgaag atgtttatga agctgttcga      900
252 acccaatccc agttggcagt gcagatccgg cggggacgag aaacactgac cttatatgtg      960
254 acccctgagg tcacagaatg a
255                                     981

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257 <210> SEQ ID NO: 4

258 <211> LENGTH: 326

259 <212> TYPE: PRT

260 <213> ORGANISM: Homo sapiens

262 <220> FEATURE:

263 <221> NAME/KEY: misc_feature

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264 <223> OTHER INFORMATION: mature HtrA2

267 <400> SEQUENCE: 4

```

269 Met Ala Val Pro Ser Pro Pro Pro Ala Ser Pro Arg Ser Gln Tyr Asn
270 1          5          10          15
273 Phe Ile Ala Asp Val Val Glu Lys Thr Ala Pro Ala Val Val Tyr Ile
274          20          25          30
277 Glu Ile Leu Asp Arg His Pro Phe Leu Gly Arg Glu Val Pro Ile Ser
278          35          40          45
281 Asn Gly Ser Gly Phe Val Val Ala Ala Asp Gly Leu Ile Val Thr Asn
282          50          55          60
285 Ala His Val Val Ala Asp Arg Arg Arg Val Arg Val Arg Leu Leu Ser
286 65          70          75          80
289 Gly Asp Thr Tyr Glu Ala Val Val Thr Ala Val Asp Pro Val Ala Asp
290          85          90          95
293 Ile Ala Thr Leu Arg Ile Gln Thr Lys Glu Pro Leu Pro Thr Leu Pro
294          100         105         110
297 Leu Gly Arg Ser Ala Asp Val Arg Gln Gly Glu Phe Val Val Ala Met
298          115         120         125
301 Gly Ser Pro Phe Ala Leu Gln Asn Thr Ile Thr Ser Gly Ile Val Ser
302          130         135         140
305 Ser Ala Gln Arg Pro Ala Arg Asp Leu Gly Leu Pro Gln Thr Asn Val
306 145         150         155         160
309 Glu Tyr Ile Gln Thr Asp Ala Ala Ile Asp Phe Gly Asn Ser Gly Gly
310          165         170         175
313 Pro Leu Val Asn Leu Asp Gly Glu Val Ile Gly Val Asn Thr Met Lys
314          180         185         190
317 Val Thr Ala Gly Ile Ser Phe Ala Ile Pro Ser Asp Arg Leu Arg Glu
318          195         200         205
321 Phe Leu His Arg Gly Glu Lys Lys Asn Ser Ser Ser Gly Ile Ser Gly
322          210         215         220
325 Ser Gln Arg Arg Tyr Ile Gly Val Met Met Leu Thr Leu Ser Pro Ser
326 225         230         235         240
329 Ile Leu Ala Glu Leu Gln Leu Arg Glu Pro Ser Phe Pro Asp Val Gln
330          245         250         255
333 His Gly Val Leu Ile His Lys Val Ile Leu Gly Ser Pro Ala His Arg
334          260         265         270
337 Ala Gly Leu Arg Pro Gly Asp Val Ile Leu Ala Ile Gly Glu Gln Met
338          275         280         285
341 Val Gln Asn Ala Glu Asp Val Tyr Glu Ala Val Arg Thr Gln Ser Gln
342          290         295         300
345 Leu Ala Val Gln Ile Arg Arg Gly Arg Glu Thr Leu Thr Leu Tyr Val
346 305         310         315         320
349 Thr Pro Glu Val Thr Glu
350          325

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353 <210> SEQ ID NO: 5

354 <211> LENGTH: 981

355 <212> TYPE: DNA

356 <213> ORGANISM: Artificial

358 <220> FEATURE:

RAW SEQUENCE LISTING

DATE: 01/10/2006

PATENT APPLICATION: US/10/562,942

TIME: 08:53:10

Input Set : A:\PTO.DA.txt

Output Set: N:\CRF4\01102006\J562942.raw

359 <223> OTHER INFORMATION: Polynucleotide consisting of the same base sequence of SEQ ID NO:

360 3 wherein the nucleotide of position 520 is g

362 <220> FEATURE:

363 <221> NAME/KEY: misc_feature

364 <223> OTHER INFORMATION: DNA that codes for mature HtrA2 (S306A)

367 <400> SEQUENCE: 5

368	atggccgctcc	ctagcccgcgc	gcccgccttct	ccccggagtc	agtacaactt	catcgagat	60
370	gtggtggaga	agacagcacc	tgccgtggtc	tatatcgaga	tcctggaccg	gcaccctttc	120
372	ttgggccg	aggtccctat	ctcgaacggc	tcaggattcg	tggtggctgc	cgatgggctc	180
374	attgtcacca	acgcccattg	ggtggctgat	cggcgagag	tccgtgtgag	actgctaagc	240
376	ggcgacacgt	atgaggccgt	ggtcacagct	gtggatccc	tggcagacat	cgcaacgctg	300
378	aggattcaga	ctaaggagcc	tctccccacg	ctgcctctgg	gacgctcagc	tgatgtccgg	360
380	caaggggagt	ttgttgttgc	catgggaagt	ccctttgcac	tgcaaacac	gatcacatcc	420
382	ggcattgtta	gctctgctca	gcgtccagcc	agagacctgg	gactccccca	aaccaatgtg	480
384	gaatacattc	aaactgatgc	agctattgat	tttggaacg	ctggagggtcc	cctggttaac	540
386	ctggatgggg	aggtgattgg	agtgaacacc	atgaagggtca	cagctggaat	ctcctttgcc	600
388	atcccttctg	atcgtcttcg	agagtttctg	catcgtgggg	aaaagaagaa	ttcctcctcc	660
390	ggaatcagt	gggtccagcg	gcgtacatt	ggggtgatga	tgctgaccct	gagtcaccgc	720
392	atccttgctg	aactacagct	tcgagaacca	agctttcccg	atgttcagca	tggtgtactc	780
394	atccataaag	tcatcctggg	ctccccctgca	caccgggctg	gtctgcggcc	tggtgatgtg	840
396	atcttgccca	ttggggagca	gatggtacaa	aatgctgaag	atgtttatga	agctgttcga	900
398	acccaatccc	agttggcagt	gcagatcccg	cggggacgag	aaacactgac	cttatatgtg	960
400	acccctgagg	tcacagaatg	a				981

403 <210> SEQ ID NO: 6

404 <211> LENGTH: 326

405 <212> TYPE: PRT

406 <213> ORGANISM: Artificial

408 <220> FEATURE:

409 <223> OTHER INFORMATION: Polypeptide consisting of the same amino acid sequence of SEQ ID

410 NO:4 wherein the 174th amino acid residue is substituted by Ala

412 <220> FEATURE:

413 <221> NAME/KEY: misc_feature

414 <223> OTHER INFORMATION: mature HtrA2 (S306A)

417 <400> SEQUENCE: 6

419	Met	Ala	Val	Pro	Ser	Pro	Pro	Pro	Ala	Ser	Pro	Arg	Ser	Gln	Tyr	Asn
420	1			5					10					15		
423	Phe	Ile	Ala	Asp	Val	Val	Glu	Lys	Thr	Ala	Pro	Ala	Val	Val	Tyr	Ile
424				20					25					30		
427	Glu	Ile	Leu	Asp	Arg	His	Pro	Phe	Leu	Gly	Arg	Glu	Val	Pro	Ile	Ser
428				35				40					45			
431	Asn	Gly	Ser	Gly	Phe	Val	Val	Ala	Ala	Asp	Gly	Leu	Ile	Val	Thr	Asn
432		50					55					60				
435	Ala	His	Val	Val	Ala	Asp	Arg	Arg	Arg	Val	Arg	Val	Arg	Leu	Leu	Ser
436	65					70					75				80	
439	Gly	Asp	Thr	Tyr	Glu	Ala	Val	Val	Thr	Ala	Val	Asp	Pro	Val	Ala	Asp
440					85				90						95	
443	Ile	Ala	Thr	Leu	Arg	Ile	Gln	Thr	Lys	Glu	Pro	Leu	Pro	Thr	Leu	Pro
444				100					105					110		
447	Leu	Gly	Arg	Ser	Ala	Asp	Val	Arg	Gln	Gly	Glu	Phe	Val	Val	Ala	Met

RAW SEQUENCE LISTING ERROR SUMMARY DATE: 01/10/2006
PATENT APPLICATION: US/10/562,942 TIME: 08:53:11

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Invalid Line Length:

The rules require that a line not exceed 72 characters in length. This includes spaces.

Seq#:1; Line(s) 6

Invalid <213> Response:

Use of "Artificial" only as "<213> Organism" response is incomplete,
per 1.823(b) of New Sequence Rules. Valid response is Artificial Sequence.

Seq#:5,6,7,8,9,10,11,12,13,14,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36

Seq#:37

VERIFICATION SUMMARY

PATENT APPLICATION: US/10/562,942

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L:11 M:270 C: Current Application Number differs, Replaced Current Application Number